

REMARKS

Claims 13-15, 31 and 32 were previously pending in this application. Claim 31 has been amended. New claim 62 has been added. As a result claims 13-15, 31-32 and 62 are pending for examination with claims 13 and 31 being independent claims. No new matter has been added.

Applicants would like to thank the Examiners for the courtesy extended in granting the telephonic interview of March 5, 2004. In the Interview Summary mailed March 8, 2004, the Examiner indicated that written description and enablement rejections were discussed. Applicant believes this is an incorrect statement of the issues discussed. Applicant agrees that the written description rejection was discussed, along with the indefiniteness rejection (35 U.S.C. §112, second paragraph). However, as there is no outstanding enablement rejection of the claims, no such rejection was discussed.

Objection to the Specification

The Examiner objected to the disclosure based on the priority information. As suggested by the Examiner, Applicant has amended the relevant paragraphs to include patent numbers. Reconsideration is respectfully requested.

Objections to the Claims

The Examiner objected to claim 31 based on several informalities. Applicant has amended the claim to limit it to the elected invention (use of antibodies in the claimed method) and to correct the text of the claim as suggested by the Examiner. Reconsideration is respectfully requested.

Rejections Under 35 U.S.C. §112, First Paragraph

The Examiner has rejected claims 13-15, 31 and 32 under 35 U.S.C. §112, first paragraph, as containing subject matter which was not described in the specification in such a way as to enable one skilled in the art to which it pertains, or with which it is most nearly connected, to make and/or use the invention. Applicant respectfully traverses the rejection.

Examiner Blanchard and Examiner Helms conducted a telephone interview on March 5, 2004 with Applicant's representatives John Van Amsterdam and Marie Jepson, and Assignee's representative Dr. Jonathan Skipper to address the issues underlying the rejection of claims 13-15, 31, and 32 under 35 U.S.C. §112. Applicants provide the following summary of the content of the telephone interview.

The interview began with a summary of the claimed invention and the related field of gastric cancer diagnostics by Dr. Jonathan Skipper. Methods set forth in the application were summarized by Dr. Skipper in the telephone interview. Dr. Skipper described the process by which the proteins that give rise to the immune response in gastric cancer were identified by Applicant using SEREX methodology. Dr. Skipper indicated that the polypeptides that Applicant identified as giving rise to an immune response in gastric cancer can be used to diagnose gastric cancer in patients. Dr. Skipper explained that the claimed methods of diagnosing gastric cancer include contacting sera obtained from a subject with sterol carrier protein-X/sterol carrier protein-2 polypeptides identified by Applicants as expressed in patients with gastric cancer. After contacting the polypeptides with the sera, one can determine whether antibodies are present in the subject's sera that recognize the polypeptide, i.e., whether the subject has mounted an immune response against the polypeptide. The presence of such antibodies in the sera is an indication of gastric cancer in that subject, either a primary tumor or a metastatic tumor that expresses the polypeptide.

Examiner Helms requested Applicants explain how the techniques used to screen patients known to have gastric cancer, as shown in the previously submitted Declaration, could be used to diagnose patients where it is not known that the patients have gastric cancer. Dr. Skipper responded stating that the patients in the study submitted in the Declaration (dated June 24, 2003, submitted July 11, 2003) were not selected, and that these results provided a demonstration of the diagnostic value of Applicant's claimed method. Applicant's representative John Van Amsterdam added that the submitted Declaration showed that normal non-cancerous patients had no sterol carrier protein-X/sterol carrier protein-2 antibodies and the technique would therefore only identify those patients with gastric cancer.

Examiner Helms requested additional information regarding the open reading frames of the identified nucleic acid sequences, SEQ ID NOs:19-22. The open reading frames for these nucleic acids are provided with this amendment (see Exhibits A-1, A-2, A-3 and A-4, translated sequences). Applicant's representative John Van Amsterdam stated that the sequences found using the SEREX method had to have open reading frames which were the same as the SCP-X protein in gastric cancer in order to be identified using this method. Dr. Skipper added that once the nucleic acid sequences are discovered they are run through a database for matches and that the identified sequences matched nucleic acid molecules encoding the sterol carrier protein-X.

Examiner Helms also requested the open reading frames because according to the Examiner, there was nothing in the claims stating the function of the SCP-X protein or how to identify which region the nucleic acid sequence matched. Dr. Skipper responded stating that it is only required that the nucleic acid encode an immunogenic protein and from knowing the nucleic acid sequence it would be routine for one of ordinary skill in the art to determine the corresponding region of SCP-X.

Examiner Blanchard asked how one would differentiate from other antibodies present in the mix. Dr. Skipper responded by providing some of the method details for a Western blot or ELISA assay for detecting an antibody of interest and stated that these were standard methods well known in the art that would distinguish between a variety of antibodies by reducing non-specific binding of antibodies.

Based on the specification as filed and on the information and arguments presented in the telephonic interview of March 5, 2004, which are summarized above, Applicant respectfully requests that the Examiner reconsider and withdraw the rejection of claims 13-15, 31, and 32 under 35 U.S.C. § 112, first paragraph.

Rejections Under 35 U.S.C. §112, Second Paragraph

The Examiner has rejected claims 31 and 32 under 35 U.S.C. §112, second paragraph, as indefinite. Applicant has amended the claims to address the first two rejections, and respectfully traverses the third rejection.

The Examiner stated three indefiniteness rejections, as follows.

(1) Claims 31 and 32 were rejected for the recitation of “parameter” in claim 31. Applicant has amended claim 31 to specify that antibodies are assayed in the claimed method. Accordingly, Applicant believes the rejection is moot.

(2) Claims 31 and 32 were rejected for the recitation in claim 31 of “said peptide of said protein,” for which there was alleged not to be antecedent basis. Applicant has amended claim 31 to remove recitations of peptides in the claimed method. Accordingly, Applicant believes this rejection is moot also.

(3) Claims 31 and 32 were rejected for the recitation of “following progress of a therapeutic regimen” in claim 31. Applicant respectfully requests reconsideration of the rejection based on the information provided in the telephonic interview.

As established, the presence of antibodies to SCP-X/SCP-2 are an indicator of gastric cancer in a patient. Patients have gastric cancer may be treated by surgery to remove the cancer, and/or radiation therapy or chemotherapy to kill the tumor, thereby removing the source of the SCP-X/SCP-2 protein. The reemergence at a later time of antibodies to SCP-X/SCP-2 would be an indication of regrowth of the tumor or of a metastatic tumor expressing SCP-X/SCP-2. Therefore, Applicant asserts that the progress of a therapeutic regime can be followed using the claimed method in which the presence or level of antibodies is determined at separate times.

In view of the foregoing claim amendments and arguments, Applicant respectfully requests reconsideration and withdrawal of the rejection of claims 31 and 32 under 35 U.S.C. §112, second paragraph.

CONCLUSION

In view of the foregoing amendments and remarks, this application should now be in condition for allowance. A notice to this effect is respectfully requested. If the Examiner believes, after this amendment, that the application is not in condition for allowance, the Examiner is requested to call the Applicant's attorney at the telephone number listed below.

If this response is not considered timely filed and if a request for an extension of time is otherwise absent, Applicant hereby requests any necessary extension of time. If there is a fee occasioned by this response, including an extension fee, that is not covered by an enclosed check, please charge any deficiency to Deposit Account No. 23/2825.

Respectfully submitted,
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Docket No. L0461.70112US00
Date: March 11, 2004
x03/11/04x

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Exhibit A-1 (SEQ ID NO: 19)

Translate Tool - Results of translation

Please select one of the following frames:

5'3' Frame 1

RQKKLF **Stop** FSIKHSSQSIILSYISLNFKK **Stop** H **Stop** Y **Stop** KN **Stop** EKR
Stop Met QIIKLT **Stop** KRKIITKD **Stop** ERYKLL **Stop** DYNLKTASESKLYCSI
 XLNDGVL **Stop** LIH **Stop** FFKKETHVKNIFILKISLCSSSHISFILIWEXN
 TVSDS **Met** KCKIFRFLISLILXTIEKLIND **Met** KCTTLITGQLLALCFLLS
 SPKGKLLN **Stop** IFSRIILKYTL **Stop** AKQKLFCLHSSLGFYCS **Stop** FYSET
 QFYPRP **Stop** LPY **Stop** LCXAQLFANS

5'3' Frame 2

ARKSYFNFLNILLKALFYPIH **Stop** ILRNNISIRKTRKKDKCR **Stop** LN
 LHEKGKL **Stop** QRTENVIN **Stop** NEIII **Stop** KLHLKANFIVQLXL **Met** **Met** VF
 YD **Stop** YTDFSRRKP **Met** LKIFLF **Stop** K **Stop** ACVQALIIFLLF **Stop** FGXKIL
 FLIA **Stop** NAKFLDF **Stop** SH **Stop** F **Stop** XLLRN **Stop** L **Met** T **Stop** SAQH **Stop** LLA
 SCWHCVSYLVLPRENS **Stop** IESSAE **Stop** SLNILCKQNKSSFFVYIVLWD
 FTVPNFILKLNFTPDHNYHINFVXHSCPLI

5'3' Frame 3

PEKVILIFY **Stop** TFFSKHYFILYLTEF **Stop** EITLVLEKLGKKINADN **Stop**
 TY **Met** KKENYNKGLRTL **Stop** IE **Met** RL **Stop** FENC **Stop** KQTLNFNYX **Stop**
Stop WCF **Met** TNTLIFQEGNPC **Stop** KYFYFKNKPVKL **Stop** SYFFYFDLG
 XKYCF **Stop** **Stop** HE **Met** QNF **Stop** IFNLTNFXNY **Stop** EID **Stop** **Stop** HEVHNTN
 YWPAVGIVFLT **Stop** FSQGKTLKLNQNNP **Stop** IYFVSKTKAFLFT
Stop FFGILLFLILF **Stop** NSILPQTIITILTLXCTVVCQF

3'5' Frame 1

Stop IGKQLCXTKLIW **Stop** LWSGVKLSFRIKLGTVKSQRT **Met** **Stop** TKKL
 LFCLQSIFKDYSIEDSI **Stop** EFSLGRTK **Stop** ETQCQQLASN **Stop** CCALH
 VINQFLNSX **Stop** N **Stop** **Stop** D **Stop** KSKNFAFHAIRNSIXFPNQNKRN **Met** I
 RA **Stop** TQAYF **Stop** NKNIFN **Met** GFLEKSVY **Stop** S **Stop** NTIIKXN **Stop** TIK
 FAFRCSFQIIISFQFITFSVLCYNFPFSCKFNYLHLSFFLVFLIL **Met** LF
 LKIQ **Stop** DIG **Stop** NNALRR **Met** FNRKLLK **Stop** LFLA

3'5' Frame 2

ELANNCAXQS **Stop** YGNYGLG **Stop** N **Stop** VSE **Stop** N **Stop** EQ **Stop** NPKELCK
 QKSFCFAYKVYLRIILLKIQFKSFPLGELSKKHANANSWPVISVVHF
Met SLINFSIVXKISEIKNLKILHF **Met** LSETVFXSQIKIKEI **Stop** SELEHR
 LIFKIKIFLTWVSFLKNQCISHKTPSLXIIIEQ **Stop** SLLSDAVFKL **Stop** SH
 FNL **Stop** RSQS FVIIFLFHVSLIICIYLFSS **Stop** FF **Stop** Y **Stop** CYFLKFSEI
Stop DKIMetL **Stop** EECLIEN **Stop** NNFFW

3'5' Frame 3

NWQTTVXNKVN**Met** VIM**Met** VWGKIEFQNKIRNSKIPKNYVNKKAFVL
LTKYI**Stop** GLFC**Stop** RFNLRVFPWEN**Stop** VRNT**Met** PTAGQ**Stop** LVLCTS
CH**Stop** SISQ**Stop** XLKLVRLKI**Stop** KFCISCYQKQYFXPKSK**Stop** KKYDQ
SLNTGLFLK**Stop** KYF**Stop** HGFPS**Stop** KISVLVIKHHH**Stop** X**Stop** LNNKV
CFQ**Met** QFSNYNLISINYVLSPLL**Stop** FSFF**Met** **Stop** V**Stop** LSAFIFFPSS
NTNVIS**Stop** NSVRYRIK**Stop** CFEKNV**Stop** **Stop** KIKITFSG

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Exhibit A-2 (SEQ ID NO: 20)

Translate Tool - Results of translation

Please select one of the following frames:

5'3' Frame 1

ILIFY **Stop** TFFSKHYFILYLTEF **Stop** XITLVLEKLGKKXNADN **Stop** TY
Met KKENYNKGLRTL **Stop** IEMet RL **Stop** FENCIS**Stop** KQTLLFNYS **Stop** **Stop**
 WCF**Met** TNTLIFQ **Stop** GNPC**Stop** KYFYFKNKPVKL **Stop** SYFFYFDLGR
 KYCF **Stop** **Stop** HEMet QNF **Stop** IFNLXNFKNY **Stop** EID **Stop** **Stop** HEVHNTN
 YWPAVGIVFLT **Stop** FSQGLLN **Stop** IFSXITLNILC **Stop** PNKTFLFT **Stop**
 FGFYCS **Stop** FYSETPFFPRP **Stop** LPYLTLLCTVV

5'3' Frame 2

F **Stop** FSIKHSSQSIILSYISLNFXX **Stop** H **Stop** Y **Stop** KN **Stop** EKRXMet QIIK
 LT **Stop** KRKIITKD **Stop** ERYKLK **Stop** DYNLKTASESKLYCSIILNDGVL
Stop LIH **Stop** FFNKETHVKNIFILKISLCSSTDHISFILIWEENTVSDS**Met**
 KCKIFRFLIXLILRTIEKLIND**Met** KCTTLITGQLLALCFLLS**SP**KENS
Stop TESSXE **Stop** P **Stop** IYFVSQTKLFCLHSSLDFTVPNFILKLHFSPDHN
 YPI **Stop** LCYAQL

5'3' Frame 3

FNFLNLILLKALFYPIH **Stop** ILXNNISIRKTRKKDXCR **Stop** LNLHEKG
 KL **Stop** QRTENVIN **Stop** NEIII **Stop** KLHLKANFIVQLFL**Met** **Met** VFYD **Stop**
 YTDFSIRKP**Met** LKIFLF **Stop** K **Stop** ACVQALIIFLLF **Stop** FGKKILFLIA
Stop NAKFLDF **Stop** SX **Stop** F **Stop** ELLRN **Stop** L**Met** T **Stop** SAQH **Stop** LLASCW
 HCVSYLVLPKTLKLNQXNNLKYTLAKQNFFVYIVLWILLFLILF
Stop NSIFPQTIITLNFV**Met** HSC

3'5' Frame 1

NNCA **Stop** QS **Stop** IG **Stop** LWSGEKWSFRIKLGTVKSKELCKQKSFVWLT
 KYI **Stop** GYXAEDSV **Stop** EFSLGELSKKHANSWPVISVVHF**Met** SLINF
 SIVLKIXEIKNLKILHF**Met** LSETVFSSQIKIKEI **Stop** SELEHRLIFKIKIF
 LTWVSLKKNQCISHKTPSLRIIEQ **Stop** SLLSDAVFKL **Stop** SHFNL **Stop** R
 SQSFVIIFLFHVSLIICIXLFS **Stop** FF **Stop** Y **Stop** CYXLKFSEI **Stop** DKIMet L
Stop EECLIEN **Stop** N


3'5' Frame 2

TTVHNKVK **Stop** GNYGLGKNGVSE **Stop** N **Stop** EQ **Stop** NPKNYVNKKVLF
 G **Stop** QSIFKVIXLKIQKSF PWEN **Stop** VRNT**Met** PTAGQ **Stop** LVLCTSC
 H **Stop** SISQ **Stop** FLKLXRLKI **Stop** KFCISCYQKQYFLPKSK **Stop** KKYDQS
 LNTGLFLK **Stop** KYF **Stop** HGFPY **Stop** KISVLVIKHHH **Stop** E **Stop** LNNKVC
 FQ**Met** QFSNYNLISYINVLSPLL **Stop** FSFF**Met** **Stop** V **Stop** LSAXIFFPSFSN
 TNVIX **Stop** NSVRYRIK **Stop** CFEKNV **Stop** **Stop** KIK

3'5' Frame 3

QLCITKLN RVIMet VWGKMet EFQNKIRNSKIQRTMet Stop TKKFCLANK
VYLRLFX Stop RFSLRVFLGR TK Stop ETQCQQLASN Stop CCALHVINQF
LNSS Stop N Stop XD Stop KSKNFAFHAI RNSIFFPNQNKRN Met IRA Stop TQ
AYF Stop NKNIFN Met GFLIEKSVY Stop S Stop NTIIKNN Stop TIKFAFRCSF
QIIISFQFITFSVLCYNFPFSCKFNYLHXSFFLVFLIL Met LFXKIQ Stop D
IG Stop NNALRR Met FNRKLLK

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Exhibit A-3 (SEQ ID NO: 21)

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Please select one of the following frames:

5'3' Frame 1

LTQLLLRCLG **Met** LEKNIWKN **Met** EQKLNTLQKLDGKIINIQLITRIPS
 SK **Met** NTV **Stop** **Met** K **Stop** WHLKKFLIF **Stop** LSYNVVPLQ **Met** VLQQQFWP
 VKHLYRS **Met** ACNPKLWKF WHKK **Stop** **Stop** LICQARLKKKALLKWLAL
 I **Stop** VKKLQENA **Met** RNLA **Stop** HQ **Met** ILT **Stop** **Stop** NFTIAFLPTNSLL **Met**
 KHWD SVQKDKVQRW LIEEIIH **Met** EESGS **Stop** ILVVD **Stop** FQRDTH **Stop**
 ALQVLLSVQNSAGS **Stop** EGKPEKRQSSWCKGGSXA **Stop** FXHWRNCG
 CNTLQDGVFPEAASSF **Stop** NSSKLKXVPTKLCKXXV **Stop** XKIXF **Stop** R
 XIEKEXXKREXGTICERKXRXEFLPFKGKXWPWGVKXGHPGVVG

5'3' Frame 2

SPSCSSDVWVCWKRTYGKIWNKN **Stop** TLCKNW **Met** EKS **Stop** TFS **Stop**
Stop PVFPVPR **Stop** IQFR **Stop** SDGI **Stop** RSF **Stop** FFDYLT **Met** LSHFRWCCS
 SNFGQ **Stop** SICTEVWPAIQSCGNFGTRNDD **Stop** FAKLV **Stop** RKKHY
Stop NGWL **Stop** YE **Stop** RSCKK **Met** L **Stop** EIWPDTK **Stop** Y **Stop** RNRTSRLLF
 YQRTPYL **Stop** STGTLSRRTRCNAG **Stop** **Stop** RR **Stop** YIWRKVGHKS **Stop**
 WWTDFKGTPTRRYRSCSVCRTLLAERGSRKRGKVPGAKVALXHN
 LXIGGTVVVTLYK **Met** GFSRKPPVPFRTHQN **Stop** SXFQPSSASXXFXX
 KSXFKXGLRRKXXRGXGEQFVKEKXXGNFCPSRGXNGPGG **Stop** KXA
 TLGWWD

5'3' Frame 3

HPVAPQ **Met** FGYAGKEH **Met** EKYGTKIEHFAKIGWKNHKHSVNNPYS
 QFQDEYSLDEV **Met** ASKEVFDLTLQCCPTSDGAAAAILASEAFVQK
 YGLQSKAVEILAQE **Met** **Met** TDLPS SFEEKSIK **Met** VGFD **Met** SKEAAR
 KCYEKSGLTPNDIDVIELHDCFSTNELLTYEALGLCPEGQGATLVDR
 GDNTYGGKWVINPSGGLISKGHPLGATGLAQCAELCWQLRGEAGK
 EAKFLVQRWLCXIIXALEELWL **Stop** HSTRWGFPGSRQFLLELIKIEA
 XSNQALQXXGLXXNXVLKXD **Stop** EGNXKEGXGNL **Stop** KKNXXGIF
 ALQGEX **Met** ALGGKRXPWGGG

3'5' Frame 1

IPPPQGGXLLPPRAIXSP **Stop** RAKIPXXFFFHKLFPXPSXXFPSQSXFK
 XRFXXKXXTCRAWLEXASIL **Met** SSKRNWRLPGKPHLVECYNHSSSN
 X **Stop** I **Met** XQSHLCTRNFASFASPLSCQQSSAH **Stop** ARPVAPSGCPFEI
 SPPLGFM **Met** THFPPYVLSPLSTSVAPCPSGQSPSAS **Stop** VRSSLVEKQS
Stop SSITSISFGVRPDFS **Stop** HFLAASLLISKPTILI **Met** LFSSNELGKSVI
 ISCAKISTALDCRPYFCTNASLAKIAAAPSEVGQHCKIVKKSKTSL
 DAITSSKLYSSWNWEYGLLTECL **Stop** FFHPIFAKCSIFVPYFSICSFPA
 YPNI **Stop** GATG **Stop**

3'5' Frame 2

SHHPRVAXFYPPGPXFPLEGQKFXPXFSTNCSXFPLXXFLLNXPLK
XDFXLNXXLAELGWNXLQF **Stop Stop** VLKGTGGFRENPI **Stop** SVTTTV
PP **Met** XKLCXRATFAPGTLPLFRLPLSAASRVLHTEQDL **Stop** RLVGVP
LKSVHH **Stop** DL **Stop** PTFH **Met** YYLLYQPALHLVLLDRVPLHK **Stop** G
VRW **Stop** KSNREVLLRQYHLVSGQISHSIFLQLLYSYQSQPF **Stop Stop** C
FFLQTSLANQSSFLVPKFPQLWIAGHTSVQ **Met** LHWPKLLLQHHLKW
DNIVR **Stop** SKNQKLL **Stop Met** PSLHLNCIHLGTGNTGY **Stop** LNVYDFSI
QFLQSVQFLFHIFPYVLFQHTQTSEEQLGE

3'5' Frame 3

PTTPGWXSFTPQGHXFPLKGKNSXXIFLSQIVPXSLFXVSFSIXL **Stop**
NXIXX **Stop** TXXLQSLVGXGFNFDEF **Stop** KELAASGKTPSCRVLQPQFL
QCXNYXAEPLHQELCLFSGFPSQLPAEFCTLSKTCSA **Stop** WVSL **Stop**
NQSTTRIYDPLSSICIISSINQRCTLSFWTESQCFISKEFVGRKAIVKF
YYVNIIWCQARFLIAFSCSFFTHIKANHFNNAFFFKRAWQISHHFLC
QNFHSFGLQAILLYKCFTGQNCCCSTI **Stop** SGTTL **Stop** DSQKIKNFFR
CHHFI **Stop** TVFILELGIRVIN **Stop Met** F **Met** IFPSNFCKVFNFCSIFFH **Met** F
FSSIPKHLRSNWV

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Exhibit A-4 (SEQ ID NO: 22)

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Please select one of the following frames:

5'3' Frame 1

G Q K X L F X F P I X X P X Q X I I X P Y X X X X X X X Y L X X **Stop** K T X E K K **Stop** X Q I
X N L X **Stop** K X K F X P X D X X X X N X X X N Y X X K X X X E T X L X G P I I X X E X X X
X L X P X F S N X E X X X K X X X I L K I T X X P T X D X I P X I W I G E K N X V X X T X X
X K X F K F L T P L X L K X X E X X I X D L N C Q P X X X A T X G X X X P Y L X P P R K X L
X X X L X Q N N P **Stop** X S L V T K X K P F X F X X S L G F N G S P I X S X T X F S P X P X L
P F Y L G K X Q X F A X P Q X S X X P X X L S P X P W E K R D X S P P **Stop** K T T F P X L W P
R X X S R L N P N N K K

5'3' Frame 2

A K K X Y X N F L L X X L X K X L F X P X X X X X X X X T F X X K K P X K K N N X K X L T
X L E X G N X Y Q X X E X X X I X X X I I X X X R X X K P X X **Met** V Q L S X X R X F X X
Stop X X D F P I X X P X L K X F X F **Stop** K **Stop** P X X Q P X I X F L X F G L G K K X X F X I
X X X A X F L N F **Stop** P P X F **Stop** X L X K X X L X T **Stop** I A N P X X X P X V G X X F L T X
S P Q G X X L X X X S X K I T L X Y P W **Stop** P X Q N L F X L X X P W D L T G P Q F X X E P
X F P X N H X Y H F T L V R X S X L X R K X V X F P X X F P P X L G K N G X G P P L K K Q
P S P X F G P X X X P V **Stop** I R T I K

5'3' Frame 3

P K K X I X I S Y **Stop** X S S X X H Y X T L X X X X F X K X P X X L K N X G K K I X A N X
Stop P X L K X E I X T X G X K X X **Stop** X E X K L X X E X G X X N Q X X W S N Y P X X G X
X X T N X X I F Q X G X X X **Stop** X X F X F K N N X X S N P X S X S F X L D W G K K X X S X
Y X E X Q X F **Stop** I F N P P X F K X Y X K X D X X L E L P T L X X G H X W X X X S L L X P
P K E X P **Stop** X E X X P K **Stop** P L X I L G N Q X K T F X V Y X X L G I **Stop** R V P N X I X N
P X F P X T I X T I L P W **Stop** G X X V C X X A X Q X X S X G X F P R X L G K T G X V P P L K
N N L P X P L A Q X X F P S K S E Q **Stop** K

3'5' Frame 1

L F I V R I **Stop** T G X X P G P K X G E G C F L R G G X I P F F P R X G G K X X G X X T X L R
X C K X L X L T K V K W **Stop** X W X G G K X G X G X N W G P V K S Q G X X K X K R F X F
G Y Q G X L R V I L X E X X X K X F P W G X **Stop** V R X X X P X G G X X X G L A I Q V X N
X X F X X F **Stop** X **Stop** G G **Stop** K F K X F X X X G X X N X I F F P N P N X R N X I X G W X
X G Y F **Stop** N X X X F X X X F X I G K X G X **Stop** X X X X L X X D N W T X K X G F X X X F
X X I X X X I X X X X V X W X K F X F S X K V X Y L X L F F F X G F L X X K V X X X X X
X X I G X N N X L X R X X N R K X K **Stop** X F L A

3'5' Frame 2

F L L F G F R R E X X L G Q R X G K V V F **Stop** G G D X S R F S Q X S G E X A X G X X L X C
X X A N X X A L P R **Stop** N G X Y G X G E X W V X D X I G D P L N P K X X V N X K G F X L
V T K D X **Stop** G L F W X X F X L R X F L G G X K **Stop** G X X X P X V A X X **Stop** G W Q F K
X X I X F X I X F K X R G V K N L K X L X F X V X G X X F F S P I Q X K G X **Stop** X G V G X

XVIFKXKXXLXXXSXLENXXISXXXPXXRIIGPXXLVXXAXFXX **Stop**
FXFXLXXFXSXGXNFXFXQXRLXICXYFFSXVF **Stop** XXRXFXKXXXX
Stop GXI**Met** XXGGXLIGNXNXFFW

3'5' Frame 3

FYCSDL DGXXXWAKXGGRLFFKGGTXPVFPKXRGKXXGEXYXFAX
XQXTXPYQGK**Met** VX**Met** VXGKXGFXXKL GTR **Stop** IPRXX **Stop** XKKVL
XWLPRIXKGYFXGXXX **Stop** XXSLGGXSKEXXAHXWXXRVGNSSX
• **Stop** XXFX **Stop** XLKXGGLKI **Stop** KXCXXXYXEXXFFPQSKXKEXDXGL
XXGLFLKXKXF **Stop** XGXXYWKIXXLXXKXPXXG **Stop** LDHXXWFXXP
XXXYNXXXNXXXFXXLVXISXFKXG **Stop** XFXIIFFXRFFXXKGXXX
XXXXXRXX **Stop** XFXEXX **Stop** **Stop** EIXIXFFG

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